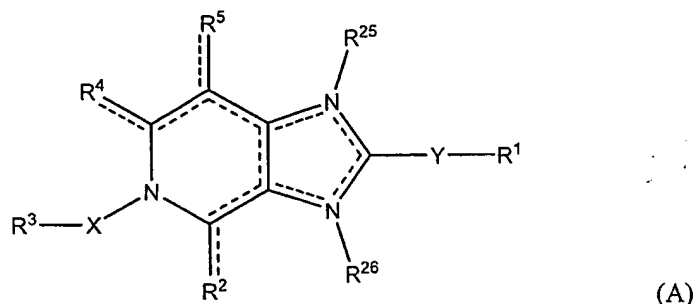


5

We claim:

1. A compound having the general formula (A),



10 wherein:

the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, optionally 4 double bonds;

R¹ is selected from hydrogen, aryl, heterocyclic, C₁-C₁₀ alkoxy,

- 15 C₁-C₁₀ thioalkyl, C₁-C₁₀ alkyl-amino, C₁-C₁₀ dialkyl-amino, C₃-₁₀ cycloalkyl, C₄-₁₀ cycloalkenyl, and C₄-₁₀ cycloalkynyl, wherein each are optionally substituted with 1 or more R⁶;

Y is selected from single bond, O, S(O)_m, NR¹¹, or C₁-₁₀ alkylene,

C₂-₁₀ alkenylene, C₂-₁₀ alkynylene, wherein each may optionally include 1 to 3

- 20 heteroatoms selected from O, S or N;

R² and R⁴ are independently selected from hydrogen, C₁-₁₈ alkyl, C₂-₁₈ alkenyl,

C₂-₁₈ alkynyl, C₁-₁₈ alkoxy, C₁-₁₈ alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸,

haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl,

- 25 C₁-₁₈ hydroxyalkyl, C₃-₁₀ cycloalkyl, C₃-₁₀ cycloalkyloxy, C₃-₁₀ cycloalkylthio, C₃-₁₀ cycloalkenyl, C₇-₁₀ cycloalkynyl, or heterocyclic, provided that when one of R²⁵ or R²⁶ is present, then either R² or R⁴ is selected from (=O), (=S), and =NR²⁷;

X is selected from C₁-C₁₀ alkylene, C₂-₁₀ alkenylene or C₂-₁₀ alkynylene, where

each may include one or more heteroatoms selected from O, S, or N, provided any such heteroatom is not adjacent to the N in the ring;

- 30 m is any integer from 0 to 2;

5 R^3 is selected from aryl, aryloxy, arylthio, cycloalkyl, cycloalkenyl, cycloalkynyl, aryl- $N(R^{10})$ -, or heterocyclic, where each said substituent may be optionally substituted with at least one R^{17} , provided that for cycloalkenyl the double bond is not adjacent to a nitrogen, and provided R^3 M-Q- is not biphenyl;

R^5 is selected from hydrogen; C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, 10 -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic;

R^6 is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halo-alkyl, C_{2-18} halo-alkenyl, C_{2-18} halo-alkynyl, C_{1-18} halo-alkoxy, C_{1-18} halo-alkylthio, C_{3-10} cycloalkyl, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, halogen, OH, CN, cyanoalkyl, -CO₂R¹⁸, NO₂, -NR⁷R⁸, C_{1-18} haloalkyl, C(=O)R¹⁸, C(=S)R¹⁸, SH, aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C_{1-18})alkyl, 15 aryl(C_{1-18})alkyloxy, aryl(C_{1-18})alkylthio, heterocyclic, C_{1-18} hydroxyalkyl, where each may be optionally substituted with at least 1 R^{19} ;

R^7 and R^8 are independently selected from hydrogen, C_{1-18} alkyl, C_{1-18} alkenyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, heterocyclic, -C(=O)R¹², -C(=S)R¹², an amino acid residue linked through a carboxyl group thereof, or where R^7 and R^8 25 together with the nitrogen form a heterocyclic;

R^9 and R^{18} are independently selected from hydrogen, OH, C_{1-18} alkyl, C_{2-18} alkenyl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, C_{1-18} alkoxy, -NR¹⁵R¹⁶, aryl, an amino acid residue linked through an amino group of the amino acid, CH₂OCH(=O)R^{9a}, or CH₂OC(=O)OR^{9a} where R^{9a} is C_1 - C_{12} alkyl, C_6 - C_{20} aryl, C_6 - C_{20} alkylaryl or C_6 - C_{20} 30 aralkyl;

R^{10} and R^{11} are independently selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, aryl, -C(=O)R¹², heterocyclic, or an amino acid residue;

R^{12} is selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, or an amino acid residue; 35

5 R^{15} and R^{16} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, or an amino acid residue;

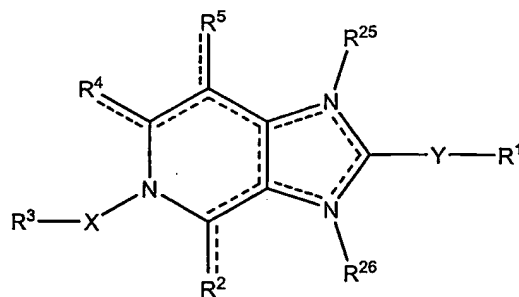
R^{17} is independently M-Q- wherein M is a ring optionally substituted with 1 or more R^{19} , and Q is a bond or a linking group connecting M to R^3 having 1 to 10 atoms and optionally substituted with 1 or more R^{19} ;

R^{19} is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{2-18} alkenyloxy, C_{2-18} alkynyloxy, C_{1-18} alkylthio, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, C_{4-10} cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹, C_{1-18} haloalkyl, C_{1-18} haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸,
 15 -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C₁₋₆ alkyl), -N(H)S(O)(O)(C₁₋₆ alkyl), aryl, heterocyclic, C_{1-18} alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C₁₋₁₈)alkyloxy, aryloxy, aryl(C₁₋₁₈ alkyl)oxy, arylthio, aryl(C₁₋₁₈)alkylthio or aryl(C₁₋₁₈)alkyl, where each may be optionally substituted with 1 or more =O, NR²⁰R²¹, CN, C_{1-18} alkoxy, heterocyclic, C_{1-18} haloalkyl, heterocyclic alkyl,
 20 heterocyclic connected to R^{17} by alkyl, alkoxyalkoxy or halogen;

R^{20} and R^{21} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, -C(=O)R¹², or -C(=S)R¹²;

R^{27} is selected from hydrogen, C_{1-18} alkyl, C_{3-10} cycloalkyl, (C₃₋₁₀ cycloalkyl)-
 25 C_{1-6} alkyl, aryl, and aryl C_{1-18} alkyl, and salts, tautomers, isomers and solvates thereof.

2. A compound having the general formula (A),



(A)

wherein:

5 the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, optionally 4 double bonds;

R^1 is selected from hydrogen, aryl, heterocyclic, C_{1-10} alkoxy, C_{1-10} thioalkyl, C_{1-10} alkyl-amino, C_{1-10} dialkyl-amino, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, and C_{4-10} cycloalkynyl, wherein each are optionally substituted with 1 or more R^6 ;

Y is selected from single bond, O, $S(O)_m$, NR^{11} , or C_{1-10} alkylene, C_{2-10} alkenylene, C_{2-10} alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O, S or N;

15 R^2 and R^4 are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkoxy, haloalkyl, -C(=O)R⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic, provided that when one of R^{25} or R^{26} is present, then either R^2 or R^4 is selected from (=O), (=S), and =NR²⁷;

X is selected from C_{1-10} alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene, where each may include one or more heteroatoms selected from O, S, or N, provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

25 R^3 is a heterocycle optionally substituted with at least one R^{17} provided, however, that R^3 optionally substituted with at least one R^{17} is not pyridinyl or 5-chlorothieryl, provided that R^3 -MQ is not biphenyl;

R^5 is selected from hydrogen; C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkoxy, haloalkyl, 30 -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic;

R^6 is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, heterocyclic, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halo-alkyl, C_{2-18} halo-alkenyl, C_{2-18} halo-alkynyl, C_{1-18} halo-alkoxy, C_{1-18} halo-alkylthio, C_{3-10} cycloalkyl, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, halogen, OH, CN,

5 cyanoalkyl, $-\text{CO}_2\text{R}^{18}$, NO_2 , $-\text{NR}^7\text{R}^8$, C_{1-18} haloalkyl, $\text{C}(=\text{O})\text{R}^{18}$, $\text{C}(=\text{S})\text{R}^{18}$, SH, aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C_{1-18})alkyl, aryl(C_{1-18})alkyloxy, aryl(C_{1-18})alkylthio, C_{1-18} hydroxyalkyl, where each may be optionally substituted with at least 1 R^{19} ;

R^7 and R^8 are independently selected from hydrogen, C_{1-18} alkyl, C_{1-18} alkenyl, 10 aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, heterocyclic, $-\text{C}(=\text{O})\text{R}^{12}$, $-\text{C}(=\text{S})\text{R}^{12}$, an amino acid residue linked through a carboxyl group thereof, or where R^7 and R^8 together with the nitrogen form a heterocyclic;

R^9 and R^{18} are independently selected from hydrogen, OH, C_{1-18} alkyl, C_{2-18} alkenyl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, C_{1-18} alkoxy, $-\text{NR}^{15}\text{R}^{16}$, aryl, an amino 15 acid residue linked through an amino group of the amino acid, $\text{CH}_2\text{OCH}(=\text{O})\text{R}^{9a}$, or $\text{CH}_2\text{OC}(=\text{O})\text{OR}^{9a}$ where R^{9a} is $\text{C}_1\text{-C}_{12}$ alkyl, $\text{C}_6\text{-C}_{20}$ aryl, $\text{C}_6\text{-C}_{20}$ alkylaryl or $\text{C}_6\text{-C}_{20}$ aralkyl;

R^{10} and R^{11} are independently selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, aryl, $-\text{C}(=\text{O})\text{R}^{12}$, 20 heterocyclic, or an amino acid residue;

R^{12} is selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, or an amino acid residue;

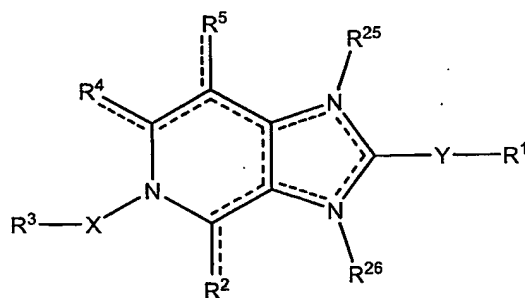
R^{15} and R^{16} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, or an amino acid 25 residue;

R^{17} is independently selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halogenated alkyl, C_{2-18} halogenated alkenyl, C_{2-18} halogenated alkynyl, C_{1-18} halogenated alkoxy, C_{1-18} halogenated alkylthio, C_{3-10} 30 cycloalkyl, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, halogen, OH, CN, CO_2H , CO_2R^{18} , NO_2 , NR^7R^8 , haloalkyl, $\text{C}(=\text{O})\text{R}^{18}$, $\text{C}(=\text{S})\text{R}^{18}$, SH, aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, arylalkyl, arylalkyloxy, arylalkylthio, heterocyclic, C_{1-18} hydroxyalkyl, where each of said aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, arylalkyl, arylalkyloxy, arylalkylthio, 35 heterocycle, or C_{1-18} hydroxyalkyl is optionally substituted with 1 or more R^{19} ;

- 5 R^{19} is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{2-18} alkenyloxy, C_{2-18} alkynyloxy, C_{1-18} alkylthio, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, C_{4-10} cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹, C_{1-18} haloalkyl, C_{1-18} haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C_{1-6} alkyl),
- 10 -N(H)S(O)(O)(C_{1-6} alkyl), aryl, heterocyclic, C_{1-18} alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C_{1-18})alkyloxy, aryloxy, aryl(C_{1-18} alkyl)oxy, arylthio, aryl(C_{1-18})alkylthio or aryl(C_{1-18})alkyl, where each may be optionally substituted with 1 or more =O, NR²⁰R²¹, CN, C_{1-18} alkoxy, heterocyclic, C_{1-18} haloalkyl, heterocyclic alkyl, heterocyclic connected to R¹⁷ by alkyl, alkoxyalkoxy or halogen;
- 15 R^{20} and R^{21} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, -C(=O)R¹², carboxylester-substituted heterocyclic or -C(=S)R¹²;
- R^{25} and R^{26} are not present, or are independently selected from hydrogen, C_{1-18} alkyl, C_{3-10} cycloalkyl, aryl, heterocyclic, where each is optionally independently
- 20 substituted with 1 to 4 of C_{1-6} alkyl, C_{1-6} alkoxy, halo, CH₂OH, benzyloxy, and OH;
- and
- R^{27} is selected from hydrogen, C_{1-18} alkyl, C_{3-10} cycloalkyl, (C_{3-10} cycloalkyl)- C_{1-6} alkyl, aryl, and aryl C_{1-18} alkyl, and
- the salts, tautomers, isomers and solvates thereof.

25

3. A compound having the general formula (A),



(A)

wherein:

5 the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, optionally 4 double bonds;

R^1 is selected from hydrogen, aryl, heterocyclic, C_1 - C_{10} alkoxy, C_1 - C_{10} thioalkyl, C_1 - C_{10} alkyl-amino, C_1 - C_{10} dialkyl-amino, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, and C_{4-10} cycloalkynyl, wherein each are optionally substituted with 1
10 or more R^6 ;

Y is selected from single bond, O, $S(O)_m$, NR^{11} , or C_{1-10} alkylene, C_{2-10} alkenylene, C_{2-10} alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O, S or N;

15 R^2 and R^4 are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic, provided that when one of R^{25} or
20 R^{26} is present, then either R^2 or R^4 is selected from (=O), (=S), and =NR²⁷;

X is selected from C_1 - C_{10} alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene, where each may include one or more heteroatoms selected from O, S, or N, provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

25 R^3 is a heterocycle optionally substituted with at least one R^{17} , provided R^3 -M-Q is not biphenyl;

R^5 is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18}
30 hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic;

R^6 is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halo-alkyl, C_{2-18} halo-alkenyl, C_{2-18} halo-alkynyl, C_{1-18} halo-alkoxy, C_{1-18} halo-alkylthio, C_{3-10}
35 cycloalkyl, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, halogen, OH, CN, cyanoalkyl, -CO₂R¹⁸, NO₂, -NR⁷R⁸, C_{1-18} haloalkyl, C(=O)R¹⁸, C(=S)R¹⁸, SH, aryl, aryloxy,

5 arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C₁₋₁₈)alkyl, aryl(C₁₋₁₈)alkyloxy, aryl(C₁₋₁₈)alkylthio, heterocyclic, C₁₋₁₈ hydroxyalkyl, where each may be optionally substituted with at least 1 R¹⁹;

R⁷ and R⁸ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₁₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, heterocyclic, -C(=O)R¹², -C(=S)R¹², an
10 amino acid residue linked through a carboxyl group thereof, or where R⁷ and R⁸ together with the nitrogen form a heterocyclic;

R⁹ and R¹⁸ are independently selected from hydrogen, OH, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₁₋₁₈ alkoxy, -NR¹⁵R¹⁶, aryl, an amino acid residue linked through an amino group of the amino acid, CH₂OCH(=O)R^{9a}, or
15 CH₂OC(=O)OR^{9a} where R^{9a} is C₁-C₁₂ alkyl, C₆-C₂₀ aryl, C₆-C₂₀ alkylaryl or C₆-C₂₀ aralkyl;

R¹⁰ and R¹¹ are independently selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, aryl, -C(=O)R¹², heterocyclic, or an amino acid residue;

20 R¹² is selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

R¹⁵ and R¹⁶ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

25 R¹⁷ is M-Q-, wherein M is a C₃₋₁₀ cycloalkyl optionally substituted with 1 or more R¹⁹, and Q is a bond, or C₁₋₁₀ alkyl optionally substituted with 1 or more R¹⁹;

R¹⁹ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₂₋₁₈ alkenyloxy, C₂₋₁₈ alkynyloxy, C₁₋₁₈ alkylthio, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₄₋₁₀ cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹,
30 C₁₋₁₈ haloalkyl, C₁₋₁₈ haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C₁₋₆ alkyl), -N(H)S(O)(O)(C₁₋₆ alkyl), aryl, heterocyclic, C₁₋₁₈alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C₁₋₁₈)alkyloxy, aryloxy, aryl(C₁₋₁₈ alkyl)oxy, arylthio, aryl(C₁₋₁₈)alkylthio or aryl(C₁₋₁₈)alkyl, where each may be optionally substituted with 1 or
35 more =O, NR²⁰R²¹, CN, C₁₋₁₈ alkoxy, heterocyclic, C₁₋₁₈ haloalkyl, heterocyclic alkyl, heterocyclic connected to R¹⁷ by alkyl, alkoxyalkoxy or halogen;

5 R^{20} and R^{21} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, $-C(=O)R^{12}$, or $-C(=S)R^{12}$;

R^{25} and R^{26} are not present, or are independently selected from hydrogen, C_{1-18} alkyl, C_{3-10} cycloalkyl, aryl, heterocyclic, where each is optionally independently
 10 substituted with 1 to 4 of C_{1-6} alkyl, C_{1-6} alkoxy, halo, CH_2OH , benzyloxy, and OH;
 and

R^{27} is selected from hydrogen, C_{1-18} alkyl, C_{3-10} cycloalkyl, $(C_{3-10}$ cycloalkyl)- C_{1-6} alkyl, aryl, and aryl C_{1-18} alkyl, and
 the salts, tautomers, isomers and solvates thereof.

15

4. The compound of claim 1, 2 or 3 wherein R^3 is heterocycle.

5. The compound of claims 1, 2 or 3 wherein YR^1 is halophenyl.

20 6. The compound of claim 5 wherein halophenyl is ortho-fluorophenyl.

7. The compound of claims 1, 2 or 3 wherein R^3 is isoxazolyl substituted with 1 R^{17} .

25 8. The compound of claims 1, 2 or 3 wherein R^{17} is aryl or an aromatic heterocycle which is substituted with 1, 2 or 3 R^{19} .

9. The compound of claims 1, 2 or 3 wherein YR^1 is none of hydrogen, an unsubstituted C_{3-10} cycloalkyl, or C_{1-6} alkyl.

30

10. The compound of claim 9 wherein YR^1 is not hydrogen.

11. The compound of claims 1, 2 or 3 wherein R^{19} is trihalomethyl, trihalomethoxy, alkoxy or halogen.

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- 5 12. The compound of claims 1, 2 or 3 wherein R¹ is aryl or aromatic heterocycle substituted with 1, 2 or 3 R⁶ wherein R⁶ is halogen, C₁₋₁₈ alkoxy; or C₁₋₁₈ haloalkyl.
13. The compound of claims 12 wherein R¹ is phenyl substituted with 1, 2 or 3 halogens.
- 10 14. The compound of claims 1, 2 or 3 wherein halogen is fluoro.
- 15 15. The compound of claims 1, 2 or 3 wherein Y is a single bond, O, C₁₋₆ alkylene, C₂₋₆ alkenylene, C₂₋₆ alkynylene or one of said groups containing 1 to 3 heteroatoms selected from O, S or NR¹¹.
16. The compound of claim 15 wherein Y is -O(CH₂)₁₋₅-, -(CH₂)₁₋₄-O-(CH₂)₁₋₄-, -S-(CH₂)₁₋₅-, -(CH₂)₁₋₄-S-(CH₂)₁₋₄-, -NR¹¹-(CH₂)₁₋₅-, -(CH₂)₁₋₄-NR¹¹-(CH₂)₁₋₄ or C₃₋₁₀ cycloalkylidene.
- 20 17. The compound of claim 15 wherein Y is -OCH₂-, -CH₂O-, C₁₋₂ alkylene, C₂₋₃ alkenylene, C₂₋₃ alkynylene, O or a bond.
18. The compound of claim 15 wherein Y is a bond.
- 25 19. The compound of claims 1, 2 or 3 wherein YR¹ is not any one of H, an unsubstituted C₃₋₁₀ cycloalkyl or C₁-C₆ alkyl.
20. The compound of claims 1, 2 or 3 wherein YR¹ is not H.
- 30 21. The compound of claims 1, 2 or 3 wherein YR¹ is halo or halomethyl-substituted phenyl.
22. The compound of claims 1, 2 or 3 wherein halo or halomethyl are ortho or meta.
- 35

- 5 23. The compound of claims 1, 2 or 3 wherein X is selected from the group consisting of alkylene, alkynylene or alkenylene and said hydrocarbons having an intrachain N, O or S heteroatom.
24. The compound of claims 1, 2 or 3 wherein X is alkyl.
- 10 25. The compound of claim 23 wherein X is selected from the group consisting of -CH₂-, -CH(CH₃)-, -CH₂-CH₂-, -CH₂-CH₂-CH₂-, -CH₂-CH₂-CH₂-CH₂-, -(CH₂)₂₋₄-O-(CH₂)₂₋₄-, -(CH₂)₂₋₄-S-(CH₂)₂₋₄-, -(CH₂)₂₋₄-NR¹⁰-(CH₂)₂₋₄-, C₃₋₁₀ cycloalkylidene, C₂₋₆ alkenylene and C₂₋₆ alkynylene.
- 15 26. The compound of claims 1, 2 or 3 wherein X is methylene.
27. The compound of claims 1, 2 or 3 wherein R³ is aryl or a heterocycle substituted with 0 to 3 R¹⁷.
- 20 28. The compound of claim 27 wherein the heterocycle is an aromatic heterocycle.
29. The compound of claim 28 wherein the heterocycle contains 1, 2 or 3 N, S or
- 25 O atoms in the ring, is linked to X through a ring carbon atom and contains 4 to 6 total ring atoms.
30. The compound of claims 1, 2 or 3 wherein R³ is isoxazolyl substituted with 1 to 3 R¹⁷.
- 30 31. The compound of claims 1, 2 or 3 wherein R¹⁷ is aryl or a heterocycle further substituted with 1 to 3 R¹⁹.
32. The compound of claims 1 or 3 wherein M is aryl or aromatic heterocycle.
- 35

- 5 33. The compound of claims 1 or 3 wherein Q contains 0 to 20 atoms selected from C, O, S, N and H.
34. The compound of claims 1 or 3 wherein M is a cyclic group selected from R¹⁷.
- 10 35. The compound of claim 2 wherein R¹⁷ is selected from the group consisting of C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, arylalkyl; arylalkyloxy; arylalkylthio; heterocycle; C₁₋₁₈ hydroxyalkyl, each of said C₃₋₁₀ cycloalkyl, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, aryl, aryloxy, arylthio, arylsulfoxide, 15 arylsulfone, arylsulfonamide, arylalkyl; arylalkyloxy; arylalkylthio; heterocycle; and C₁₋₁₈ hydroxyalkyl is unsubstituted or is substituted 1 or more R¹⁹.
36. The compound of claim 2 wherein R¹⁷ is selected from the group consisting of aryl and heterocycle, and where said aryl or heterocycle is optionally substituted 20 with 1 or more R¹⁹.
37. The compound of claims 1, 2 or 3 wherein R⁹ and R¹⁸ are H, OH or alkyl.
38. The compound of claims 1, 2 or 3 wherein R⁵ is H.
- 25 39. The compound of claims 1, 2 or 3 wherein R⁶ is halogen.
40. The compound of claims 1, 2 or 3 wherein R⁷, R⁸, R¹⁰, R¹¹, R¹⁵, R¹⁶, R²⁰, and R²¹ are independently H or C₁₋₁₈ alkyl.
- 30 41. The compound of claims 1, 2 or 3 wherein R¹² is OH or alkyl.
42. The compound of claims 1, 2 or 3 wherein R¹⁹ is selected from the group consisting of H; C₁₋₁₈ alkyl; C₂₋₁₈ alkenyl; C₂₋₁₈ alkynyl; C₁₋₁₈ alkoxy; alkenyloxy; 35 alkynyloxy; C₁₋₁₈ alkylthio; C₃₋₁₀ cycloalkyl; C₄₋₁₀ cycloalkenyl; C₄₋₁₀ cycloalkynyl; halogen; OH; CN; cyanoalkyl; NO₂; NR²⁰R²¹; haloalkyl; haloalkyloxy; C(=O)R¹⁸;

- 5 C(=O)OR¹⁸; OalkenylC(=O)OR¹⁸; -OalkylC(=O)NR²⁰R²¹; aryl; heterocycle; -
 OalkylOC(=O)R¹⁸; C(=O)N(C₁₋₆ alkyl), N(H)S(O)(O)(C₁₋₆ alkyl); arylalkyloxy;
 aryloxy; arylalkyloxy; and arylalkyl; each of which is unsubstituted or substituted
 with 1 or more =O; NR²⁰R²¹; CN; alkoxy; heterocycle; haloalkyl- or alkyl-
 substituted heterocycle; and heterocycle linked to R¹⁷ by alkyl; alkoxyalkoxy or
 10 halogen.

43. The compound of claim 42 wherein R¹⁹ is independently selected from the
 group consisting of halogen, N(R²⁰ R²¹), alkoxy, halo-substituted alkyl and halo-
 substituted alkoxy.

15

44. The compound of claims 1, 2 or 3 wherein R²⁵ and R²⁶ are not present.

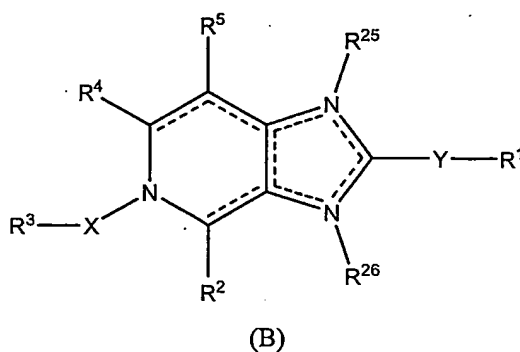
45. The compound of claims 1, 2 or 3 which is not substituted at R²⁵ but is
 substituted at R²⁶, and either R² or R⁴ is selected from (=O), (=S), and (=NR²⁷).

20

46. The compound of claims 1, 2 or 3 wherein haloalkyl or haloalkyloxy is -CF₃
 or -OCF₃.

47. A composition comprising a pharmaceutically acceptable excipient and a
 25 compound of claims 1, 2 or 3.

48. A compound having the general formula (B),



30

wherein:

5 the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3, optionally 4 double bonds;

R^1 is selected from hydrogen, aryl, heterocyclic, C_{1-10} alkoxy, C_{1-10} thioalkyl, C_{1-10} alkyl-amino, C_{1-10} dialkyl-amino, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, and C_{4-10} cycloalkynyl, wherein each are optionally substituted with 1 or more R^6 ;

Y is selected from single bond, O, $S(O)_m$, NR^{11} , or C_{1-10} alkylene, C_{2-10} alkenylene, C_{2-10} alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O, S or N;

15 R^2 and R^4 are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic, provided that when one of R^{25} or R^{26} is present, then either R^2 or R^4 is selected from (=O), (=S), and =NR²⁷;

X is selected from C_{1-10} alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene, where each may include one or more heteroatoms selected from O, S, or N, provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

25 R^3 is selected from aryl, aryloxy, arylthio, cycloalkyl, cycloalkenyl, cycloalkynyl, aryl-N(R¹⁰)-, or heterocyclic, where each said substituent may be optionally substituted with at least one R^{17} , provided that for cycloalkenyl the double bond is not adjacent to a nitrogen, and provided R^3 M-Q- is not biphenyl;

30 R^5 is selected from hydrogen; C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic;

35 R^6 is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halo-alkyl, C_{2-18} halo-alkenyl, C_{2-18} halo-alkynyl, C_{1-18} halo-alkoxy, C_{1-18} halo-alkylthio, C_{3-10}

5 cycloalkyl, C₃₋₁₀ cycloalkenyl, C₇₋₁₀ cycloalkynyl, halogen, OH, CN, cyanoalkyl, -CO₂R¹⁸, NO₂, -NR⁷R⁸, C₁₋₁₈ haloalkyl, C(=O)R¹⁸, C(=S)R¹⁸, SH, aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C₁₋₁₈)alkyl, aryl(C₁₋₁₈)alkyloxy, aryl(C₁₋₁₈)alkylthio, heterocyclic, C₁₋₁₈ hydroxyalkyl, where each may be optionally substituted with at least 1 R¹⁹;

10 R⁷ and R⁸ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₁₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, heterocyclic, -C(=O)R¹², -C(=S)R¹², an amino acid residue linked through a carboxyl group thereof, or where R⁷ and R⁸ together with the nitrogen form a heterocyclic;

R⁹ and R¹⁸ are independently selected from hydrogen, OH, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₁₋₁₈ alkoxy, -NR¹⁵R¹⁶, aryl, an amino acid residue linked through an amino group of the amino acid, CH₂OCH(=O)R^{9a}, or CH₂OC(=O)OR^{9a} where R^{9a} is C₁-C₁₂ alkyl, C₆-C₂₀ aryl, C₆-C₂₀ alkylaryl or C₆-C₂₀ aralkyl;

R¹⁰ and R¹¹ are independently selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, aryl, -C(=O)R¹², heterocyclic, or an amino acid residue;

R¹² is selected from the group consisting of hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

R¹⁵ and R¹⁶ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, or an amino acid residue;

R¹⁷ is independently M-Q- wherein M is a ring optionally substituted with 1 or more R¹⁹, and Q is a bond or a linking group connecting M to R³ having 1 to 10 atoms and optionally substituted with 1 or more R¹⁹;

30 R¹⁹ is selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, C₁₋₁₈ alkoxy, C₂₋₁₈ alkenyloxy, C₂₋₁₈ alkynyloxy, C₁₋₁₈ alkylthio, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, C₄₋₁₀ cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹, C₁₋₁₈ haloalkyl, C₁₋₁₈ haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkyloC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C₁₋₆ alkyl),
35 -N(H)S(O)(O)(C₁₋₆ alkyl), aryl, heterocyclic, C₁₋₁₈alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C₁₋₁₈)alkyloxy, aryloxy, aryl(C₁₋₁₈ alkyl)oxy, arylthio, aryl(C₁₋

5 ₁₈)alkylthio or aryl(C₁₋₁₈)alkyl, where each may be optionally substituted with 1 or more =O, NR²⁰R²¹, CN, C₁₋₁₈ alkoxy, heterocyclic, C₁₋₁₈ haloalkyl, heterocyclic alkyl, heterocyclic connected to R¹⁷ by alkyl, alkoxyalkoxy or halogen;

R²⁰ and R²¹ are independently selected from hydrogen, C₁₋₁₈ alkyl, C₂₋₁₈ alkenyl, C₂₋₁₈ alkynyl, aryl, C₃₋₁₀ cycloalkyl, C₄₋₁₀ cycloalkenyl, -C(=O)R¹², or
 10 -C(=S)R¹²;

R²⁷ is selected from hydrogen, C₁₋₁₈ alkyl, C₃₋₁₀ cycloalkyl, (C₃₋₁₀ cycloalkyl)-C₁₋₆ alkyl, aryl, and aryl C₁₋₁₈ alkyl, and salts, tautomers, isomers and solvates thereof.

49. The compound of claim 48 wherein Y is a single bond, and R¹ is aryl.

15

50. The compound of claim 48 wherein X is C₁-C₁₀ alkylene, C₂₋₁₀ alkenylene or C₂₋₁₀ alkynylene.

51. The compound of claim 48 wherein R³ is heterocyclic.

20

52. The compound of claim 48 wherein R³ is heterocyclic substituted with R¹⁷ where Q is a bond and M is aryl.

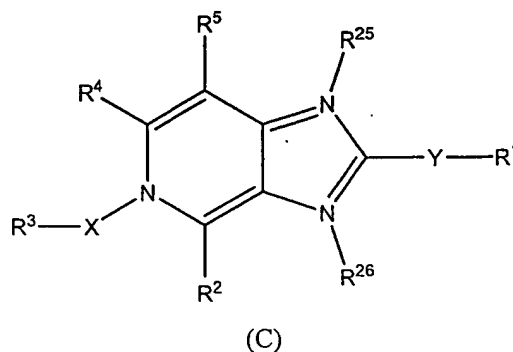
53. The compound of claim 48 wherein Y is a single bond, and R¹ is phenyl.

25

54. The compound of claim 48 wherein R³ is isoxazole substituted with R¹⁷ where Q is a bond and M is aryl.

55. The compound of claim 48 wherein R³ is isoxazole substituted with R¹⁷ where
 30 Q is a bond and M is phenyl.

56. A compound having the general formula (C),



5

wherein:

the dotted lines represent an optional double bond, provided that no two double bonds are adjacent to one another, and that the dotted lines represent at least 3,
10 optionally 4 double bonds;

R^1 is selected from hydrogen, aryl, heterocyclic, C_1 - C_{10} alkoxy, C_1 - C_{10} thioalkyl, C_1 - C_{10} alkyl-amino, C_1 - C_{10} dialkyl-amino, C_3 - C_{10} cycloalkyl, C_4 - C_{10} cycloalkenyl, and C_4 - C_{10} cycloalkynyl, wherein each are optionally substituted with 1 or more R^6 ;

15 Y is selected from single bond, O, $S(O)_m$, NR^{11} , or C_1 - C_{10} alkylene, C_2 - C_{10} alkenylene, C_2 - C_{10} alkynylene, wherein each may optionally include 1 to 3 heteroatoms selected from O, S or N;

R^2 and R^4 are independently selected from hydrogen, C_1 - C_{18} alkyl, C_2 - C_{18} alkenyl, C_2 - C_{18} alkynyl, C_1 - C_{18} alkoxy, C_1 - C_{18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸,
20 haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_1 - C_{18} hydroxyalkyl, C_3 - C_{10} cycloalkyl, C_3 - C_{10} cycloalkyloxy, C_3 - C_{10} cycloalkylthio, C_3 - C_{10} cycloalkenyl, C_7 - C_{10} cycloalkynyl, or heterocyclic, provided that when one of R^{25} or R^{26} is present, then either R^2 or R^4 is selected from (=O), (=S), and =NR²⁷;

X is selected from C_1 - C_{10} alkylene, C_2 - C_{10} alkenylene or C_2 - C_{10} alkynylene, where
25 each may include one or more heteroatoms selected from O, S, or N, provided any such heteroatom is not adjacent to the N in the ring;

m is any integer from 0 to 2;

R^3 is selected from aryl, aryloxy, arylthio, cycloalkyl, cycloalkenyl, cycloalkynyl, aryl-N(R^{10})-, or heterocyclic, where each said substituent may be
30 optionally substituted with at least one R^{17} , provided that for cycloalkenyl the double bond is not adjacent to a nitrogen, and provided R^3 M-Q- is not biphenyl;

- 5 R^5 is selected from hydrogen; C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, halogen, -OH, -CN, -NO₂, -NR⁷R⁸, haloalkyloxy, haloalkyl, -C(=O)R⁹, -C(=O)OR⁹, -C(=S)R⁹, SH, aryl, aryloxy, arylthio, arylalkyl, C_{1-18} hydroxyalkyl, C_{3-10} cycloalkyl, C_{3-10} cycloalkyloxy, C_{3-10} cycloalkylthio, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, or heterocyclic;
- 10 R^6 is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{1-18} alkylthio, C_{1-18} alkylsulfoxide, C_{1-18} alkylsulfone, C_{1-18} halo-alkyl, C_{2-18} halo-alkenyl, C_{2-18} halo-alkynyl, C_{1-18} halo-alkoxy, C_{1-18} halo-alkylthio, C_{3-10} cycloalkyl, C_{3-10} cycloalkenyl, C_{7-10} cycloalkynyl, halogen, OH, CN, cyanoalkyl, -CO₂R¹⁸, NO₂, -NR⁷R⁸, C_{1-18} haloalkyl, C(=O)R¹⁸, C(=S)R¹⁸, SH, aryl, aryloxy, arylthio, arylsulfoxide, arylsulfone, arylsulfonamide, aryl(C_{1-18})alkyl, aryl(C_{1-18})alkyloxy, aryl(C_{1-18})alkylthio, heterocyclic, C_{1-18} hydroxyalkyl, where each may be optionally substituted with at least 1 R¹⁹;
- R^7 and R^8 are independently selected from hydrogen, C_{1-18} alkyl, C_{1-18} alkenyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, heterocyclic, -C(=O)R¹², -C(=S)R¹², an amino acid residue linked through a carboxyl group thereof, or where R^7 and R^8 together with the nitrogen form a heterocyclic;
- 20 R^9 and R^{18} are independently selected from hydrogen, OH, C_{1-18} alkyl, C_{2-18} alkenyl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, C_{1-18} alkoxy, -NR¹⁵R¹⁶, aryl, an amino acid residue linked through an amino group of the amino acid, CH₂OCH(=O)R^{9a}, or
- 25 CH₂OC(=O)OR^{9a} where R^{9a} is C_1 - C_{12} alkyl, C_6 - C_{20} aryl, C_6 - C_{20} alkylaryl or C_6 - C_{20} aralkyl;
- R^{10} and R^{11} are independently selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, aryl, -C(=O)R¹², heterocyclic, or an amino acid residue;
- 30 R^{12} is selected from the group consisting of hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, or an amino acid residue;
- R^{15} and R^{16} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, or an amino acid residue;

5 R^{17} is independently M-Q- wherein M is a ring optionally substituted with 1 or more R^{19} , and Q is a bond or a linking group connecting M to R^3 having 1 to 10 atoms and optionally substituted with 1 or more R^{19} ;

R^{19} is selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, C_{1-18} alkoxy, C_{2-18} alkenyloxy, C_{2-18} alkynyloxy, C_{1-18} alkylthio, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, C_{4-10} cycloalkynyl, halogen, -OH, -CN, cyanoalkyl, -NO₂, -NR²⁰R²¹, C_{1-18} haloalkyl, C_{1-18} haloalkyloxy, -C(=O)R¹⁸, -C(=O)OR¹⁸, -OalkenylC(=O)OR¹⁸, -OalkylC(=O)NR²⁰R²¹, -OalkylOC(=O)R¹⁸, -C(=S)R¹⁸, SH, -C(=O)N(C₁₋₆ alkyl), -N(H)S(O)(O)(C₁₋₆ alkyl), aryl, heterocyclic, C_{1-18} alkylsulfone, arylsulfoxide, arylsulfonamide, aryl(C₁₋₁₈)alkyloxy, aryloxy, aryl(C₁₋₁₈ alkyl)oxy, arylthio, aryl(C₁₋₁₈)alkylthio or aryl(C₁₋₁₈)alkyl, where each may be optionally substituted with 1 or more =O, NR²⁰R²¹, CN, C_{1-18} alkoxy, heterocyclic, C_{1-18} haloalkyl, heterocyclic alkyl, heterocyclic connected to R^{17} by alkyl, alkoxyalkoxy or halogen;

R^{20} and R^{21} are independently selected from hydrogen, C_{1-18} alkyl, C_{2-18} alkenyl, C_{2-18} alkynyl, aryl, C_{3-10} cycloalkyl, C_{4-10} cycloalkenyl, -C(=O)R¹², or -C(=S)R¹²;

R^{27} is selected from hydrogen, C_{1-18} alkyl, C_{3-10} cycloalkyl, (C₃₋₁₀ cycloalkyl)-C₁₋₆ alkyl, aryl, and aryl C₁₋₁₈ alkyl, and salts, tautomers, isomers and solvates thereof.

25 57. The compound of claim 56 wherein Y is a single bond, and R^1 is aryl.

58. The compound of claim 56 wherein X is C_1 - C_{10} alkylene, C_{2-10} alkenylene or C_{2-10} alkynylene.

30 59. The compound of claim 56 wherein R^3 is heterocyclic.

60. The compound of claim 56 wherein R^3 is heterocyclic substituted with R^{17} where Q is a bond and M is aryl.

35 61. The compound of claim 56 wherein Y is a single bond, and R^1 is phenyl.

5 62. The compound of claim 56 wherein R³ is isoxazole substituted with R¹⁷ where Q is a bond and M is aryl.

63. The compound of claim 56 wherein R³ is isoxazole substituted with R¹⁷ where Q is a bond and M is phenyl.

10

64. A method comprising administering to a subject in need of treatment or prophylaxis of a viral infection an antivirally effective amount of a compound of claims 1, 2, 3, 48 or 56.

15 65. The method of claim 64, wherein the viral infection is an infection of a hepatitis-C virus.

66. The method of claim 65 further comprising administering at least one additional antiviral therapy to the subject.

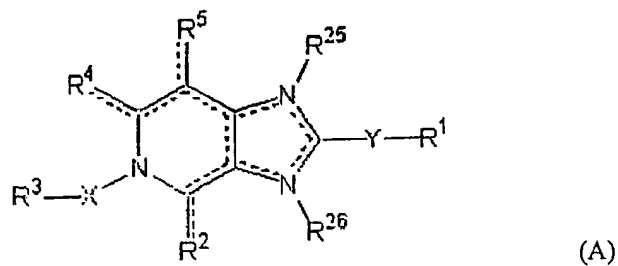
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67. The method of claim 66 wherein the additional therapy is selected from the group consisting of an interferon alpha and ribavirin.

25 68. A method of screening antiviral compounds which comprises providing a compound of claims 1, 2, 3, 48 or 56 and determining the anti-viral activity of said compound.

69. The method of claim 68 wherein said anti-viral activity is determined by the activity of said compound against one or more viruses belonging to the family of the
30 Flaviviridae and/or of the Picornaviridae.

70. A method for assaying the structure-activity of analogues of formula (A) compounds



wherein the substituents are defined in WO 2004/005286, comprising

- (c) preparing a compound of formula (A) in which at least one substituent is not disclosed by WO 2004/005286; and
- 10 (d) determining the anti-HCV activity of the compound of step (a).

71. The method of claim 70 wherein the substituent is located at R^3 , R^2 , R^4 , R^{26} and/or R^5 .

15